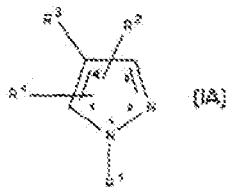




INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

<p>(51) International Patent Classification ⁷ :</p> <p>C07D 401/04, A61K 31/415, 31/47, 31/445, 31/44, 31/50, 31/505, 31/52, C07D 405/14, 401/14, 409/14, 403/04, 487/04, 473/00, 413/14, 417/14, 471/04, A61P 29/00 // (C07D 487/04, 293:00, 231:00) (C07D 471/04, 221:00, 209:00)</p>	<p>A1</p>	<p>(11) International Publication Number: WO 00/31063</p> <p>(43) International Publication Date: 2 June 2000 (02.06.00)</p>
<p>(21) International Application Number: PCT/US99/26007</p> <p>(22) International Filing Date: 17 November 1999 (17.11.99)</p> <p>(30) Priority Data: 09/196,623 20 November 1998 (20.11.98) US</p> <p>(71) Applicant (for all designated States except US): G.D. SEARLE & CO. [US/US]; Corporate Patent Dept., P.O. Box 5110, Chicago, IL 60680-5110 (US).</p> <p>(72) Inventors; and</p> <p>(75) Inventors/Applicants (for US only): AMANTANARAYAN, Ashok [US/US]; 54 Lisk Drive, Hainesville, IL 60030 (US); CLARE, Michael [GB/US]; 5154 West Brown Street, Skokie, IL 60077 (US); COLLINS, Paul W. [US/US]; 2317 Carillon Drive, Grayslake, IL 60030 (US); CRICH, Joyce, Z. [US/US]; 1501 G. Topp Lane, Glenview, IL 60025 (US); DEVRAJ, Rajesh [IN/US]; 1331 Oak Borough Drive, Ballwin, MO 63021 (US); FLYNN, Daniel L. [US/US]; One Axygen Center Drive, Thousand Oaks, CA 91320 (US); GENG, Lifeng [CN/US]; 5300 Davis Street, Skokie, IL 60077 (US); GRANETO, Matthew, J. [US/US]; 352 Hartwell Court, Chesterfield, MO 63017 (US); HANAU, Cathleen, E. [US/US]; 15551 Wendimill Drive, Chesterfield, MO 63017 (US); HANSON, Gunnar, J. [US/US]; 7410 Keystone Avenue, Skokie, IL 60077 (US); HARTMANN, Susan, J. [US/US]; 513 West Essex Avenue, Kirkwood, MO 63122 (US); HEPPELLE, Michael [DE/US]; 1460 Keystone Court, St. Charles, MO 63303 (US); HUANG, He [CN/US]; 2309F S. Stewart Avenue, Chicago, IL 60616 (US); KHANNA, Ish, K. [CN/US]; 1821 S. Pulcon Drive, Libertyville, IL 60048 (US); KOSZYK, Francis, J. [US/US]; 11 Wildwood Drive South, Prospect Heights,</p>		<p>IL 60070 (US); LIAO, Shuyuan [US/US]; 1536 Wabers Avenue, Northbrook, IL 60062 (US); METZ, Suzanne [US/US]; 525 Westernmill Drive, Chesterfield, MO 63017 (US); PARTIS, Richard, A. [US/US]; 2221 Noyes Street, Evanston, IL 60201 (US); PERRY, Thao, D. [US/US]; 516 Bloom Street, Red Bud, IL 62278 (US); RAO, Shashidhar, N. [IN/US]; 736 Bellerive Manor Drive, St. Louis, MO 63141 (US); SELNESS, Shaun, Raj [US/US]; 1875 Cedarhill Drive, Chesterfield, MO 63017 (US); SOUTH, Michael, S. [US/US]; 11671 Cheftain Drive, Chesterfield, MO 63146 (US); STEALEY, Michael, A. [US/US]; 502 Juniper Parkway, Libertyville, IL 60048 (US); TALLEY, John, Jeffrey [US/US]; 4701 Clifton Avenue, St. Louis, MO 63109 (US); VAZQUEZ, Michael, L. [US/US]; 614 Castle Meadows Court, Ballwin, MO 63021 (US); WEBER, Richard, M. [US/US]; 240 Hickory Court, Lake Bluff, IL 60044 (US); XU, Xiangdong [CN/US]; 7498 Abbey Road, Gurnee, IL 60031 (US); YU, Yi [CN/US]; 9065 Grasse Point Road, Skokie, IL 60077 (US).</p> <p>(74) Agents: SCRIVNER, Alan, L. et al.; G.D. Searle & Co., Corporate Patent Dept., P.O. Box 5110, Chicago, IL 60680-5110 (US).</p> <p>(81) Designated States: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).</p> <p>Published With international search report. Before the expiration of the time limit for amending the claims and to be republished in the event of the receipt of amendments.</p>

(54) Title: SUBSTITUTED PYRAZOLES AS p38 KINASE INHIBITORS



(57) Abstract

A class of pyrazole derivatives is described for use in treating p38 kinase mediated disorders. Compounds of particular interest are defined by Formula (IA), wherein R¹, R², R³ and R⁴ are as described in the specification.